

Serial No.: 08/270,412  
Filed: July 5, 1994

pursuant to 37 C.F.R. §1.132 (and attached Appendices A-D), and Exhibits A-E. Original declarations will be provided at the Examiner's request.

Kindly amend the present application in the following respects:

In the Claims:

17. (Twice Amended) A method for the *in vitro* proliferation of a multipotent neural stem cell comprising the steps of:

(a) dissociating mammalian neural tissue containing at least one multipotent stem cell capable of producing progeny that are capable of differentiating into neurons and glia; and

(b) exposing said [dissociated] multipotent stem cell to a first culture medium containing at least one growth factor to produce progeny of said multipotent stem cell, and

(c) passaging said progeny to a second culture medium containing at least one growth factor to proliferate said progeny [cell].

In Claim 85, after "amphiregulin," insert --fibroblast growth factor--.

In Claim 87, delete "preexposed" and replace with --exposed--.

Sub E2 → 88. (Amended) The method of Claim 17 wherein step (c) is repeated at least one additional time[said multipotent stem cell is capable of proliferation *in vitro* without limit].

89. (Amended) The method of Claim 17 wherein the progeny passaged in step (c) are [said cell is proliferated] in suspension.

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*Such D2*  
90. (Amended) The method of Claim 17 further comprising the additional step of:

[(c)] (d) inducing the progeny [cells] proliferated in step [(b)] (c) to differentiate by plating said [cells] progeny on a fixed substrate.

91. (Amended) The method of Claim 17 further comprising the additional step of:

*C2 Cont*  
[(c)] (d) inducing the progeny [cells] proliferated in step [(b)] (c) to differentiate in suspension by allowing [the cells] said progeny to form [clusters of cells] clonally-derived neurospheres without reinitiating proliferation.

*Such D3*  
93. (Amended) The method of Claim 17 wherein the progeny produced in step (b) grow in the form of a clonally-derived neurosphere [said dissociated multipotent stem cell is proliferated at least 21 days *in vitro* with substantially no differentiation].

Please add the following claim:

*C Such D2*  
--94. The method of Claim 93 wherein the passaging of said progeny in step (c) is achieved by dissociating said neurosphere to form a suspension of single cells and suspending said cells in said second culture medium.--

#### REMARKS

Please charge any additional fees or credit overpayments to Deposit Account No. 06-1300 (Order No. A-57660-1/DJB). A duplicate copy of this sheet is enclosed.